

In the claims:

This listing of claims will replace all prior versions, and listings, of claims in the application.

1-21. (cancelled)

22. (currently amended) A method for designing proteins with previously unknown, realizable backbone configurations comprising:

generating backbone protein configurations of ~~fixed~~ a preselected length using a set of dihedral angle pairs;

assigning a sphere of fixed radius or another space-filling generic side chain to each position where an amino acid residue would ultimately be inserted in the generated configurations;

eliminating self-intersecting configurations;

evaluating ~~the~~ a surface exposure of each sphere or generic side chain in each remaining configuration;

normalizing the total surface exposure of each remaining configuration;

generating sequences of hydrophobicities having the same length as the number of spheres or generic side chains in each of the remaining generated configurations;

determining, for each sequence of hydrophobicities, which of the remaining configurations is the ground state;

recording the ground-state configuration for each sequence considered;

~~identifying as highly designable~~ selecting those configurations which are both ground states of the largest number of sequences and novel; and

selecting novel sequences of amino acids designed to adopt one of the ~~highly designable~~ selected configurations.

23. (original) A method for designing proteins as in claim 22 wherein:

one set of dihedral angle pairs corresponds to an alpha helix and one set of dihedral angles corresponds to a beta strand.

24. (original) A method for designing proteins as in claim 22 wherein:

two sets of dihedral angles correspond to an alpha helix and one set of dihedral angle pairs corresponds to a beta strand.

25. (original) A method for designing proteins as in claim 24 wherein:

additional dihedral angle pairs fall within regions of high frequency in a Ramachandran plot.

26. (previously presented) A method for designing proteins as in claim 22 wherein:

the probability of choosing a particular pair of dihedral angles depends on the preceding pairs of dihedral angles along the backbone.

27. (cancelled)

28. (previously presented) A method for designing proteins as in claim 22 further comprising:
eliminating non-compact configurations.

29. (previously presented) A method for designing proteins as in claim 28 further comprising:

clustering configurations which are sufficiently similar in the three dimensional trajectories of their backbones and considering all configurations within such a cluster to be variants of a single configuration;

summing, for all configurations in a cluster, the number of sequences with that configuration as their ground state; and identifying as highly designable those clusters of configurations with the largest sum of associated sequences.

30. (cancelled)

31. (original) A method for designing proteins as in claim 22 wherein:

the set of dihedral angles is a set of strings of dihedral angles.

32. (original) A method for designing proteins as in claim 31 wherein:

the strings of angles are weighted according to their frequency of appearance in natural proteins and infrequent strings are eliminated.

33. (previously presented) A method for designing proteins as in claim 22 wherein:

normalizing is accomplished by dividing the surface exposure of each sphere or generic side chain assigned to a given configuration by the total surface exposure of that configuration.

34. (cancelled)

35. (original) A method for designing proteins as in claim 22 further comprising:

eliminating non-compact configurations after self-intersecting configurations are eliminated.

36. (previously presented) A method for designing proteins as in claim 35 further comprising:

clustering configurations which are sufficiently similar in the three dimensional trajectories of their

backbones and considering all configurations within a cluster to be variants of a single configuration;

summing, for all configurations in a cluster, the number of sequences with that configuration as their ground state; and identifying as highly designable those clusters of configurations with the largest sum of associated sequences.

37. (original) A method for designing proteins as in claim 22 further comprising:

eliminating all configurations that are not favorable for forming hydrogen bonds after eliminating non-compact configurations.

38. (previously presented) A method for designing proteins as in claim 22 further comprising:

clustering configurations which are sufficiently similar in the three dimensional trajectories of their backbones and considering all configurations within a cluster to be variants of a single configuration;

summing, for all configurations in a cluster, the number of sequences with that configuration as their ground state; and identifying as highly designable those clusters of configurations with the largest sum of associated sequences.

39. (original) A method for designing proteins as in claim 38 wherein:

clustering is accomplished by totaling the root-mean-square distance between every pair of configurations and defining a configuration as a member of a cluster if it lies within a root-mean-square distance λ of any member of the cluster.

40. (original) A method for designing proteins as in claim 39 wherein:

λ is 0.4 Angstroms per amino acid.

41-57. (cancelled)